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AMENDMENT AND RESPONSE TO RESTRICTION REQUIREMENT

Amendment

In the Claims

- 1. (original) A compound comprising a target specific portion and an effector portion wherein:
- (i) the target specific portion comprises or consists of a monoclonal antibody having specificity for oncofoetal fibronectin, or a fragment or variant thereof which retains the binding specificity for oncofoetal fibronectin of the parent monoclonal antibody; and
- (ii) the effector portion comprises or consists of interleukin-12, or a functional fragment or variant thereof

characterised in the monoclonal antibody having specificity for oncofoetal fibronectin binds to a region of oncofoetal fibronectin other than the ED-B region.

- 2. (original) A compound according to Claim 1 wherein the target specific portion is capable of binding to an amino acid sequence within the repeat 7 domain of fibronectin.
- 3. (previously presented) A compound according to Claim 1 wherein the target specific portion is capable of binding an amino acid sequence within the repeat 7 domain of fibronectin.
- 4. (previously presented) A compound according to Claim 1 wherein the target specific portion is specific for human oncofoetal fibronectin.
- 5. (previously presented) A compound according to Claim 1 wherein the monoclonal antibody having specificity for oncofoetal fibronectin is a BC1 antibody, or an antibody capable of competing with the binding of a BC1 antibody to oncofoetal fibronectin.

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- 6. (original) A compound according to Claim 5 wherein the monoclonal antibody having specificity for oncofoetal fibronectin is a BC1 antibody.
- 7. (previously presented) A compound according to Claim 1 wherein the monoclonal antibody is a human or humanized antibody.
- 8. (previously presented) A compound according to Claim 6 wherein the compound binds to oncofoetal fibronectin more tightly than the parent monoclonal antibody.
- 9. (original) A compound according to Claim 8 wherein the compound binds to oncofoetal fibronectin more at least 2-fold tighter than the parent monoclonal antibody.
- 10. (previously presented) A compound according to Claim 8 wherein the compound binds to oncofoetal fibronectin at least 10-fold tighter than the parent BC1 antibody binds to oncofoetal fibronectin.
- 11. (previously presented) A compound according to Claim 1 wherein the target specific portion comprises a polypeptide of SEQ ID NO: 1.
- 12. (previously presented) A compound according to Claim 1 wherein the target specific portion comprises a polypeptide of SEQ ID NO: 2.
- 13. (previously presented) A compound according to Claim 11 wherein the target specific portion comprises a polypeptide of SEQ ID NO: 1 and a polypeptide SEQ ID NO: 2.
- 14. (previously presented) A compound according to Claim 1 wherein the target specific portion comprises or consists of an antigen binding fragment of a monoclonal antibody having specificity for oncofoetal fibronectin.

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- 15. (original) A compound according to Claim 14 wherein the target specific portion comprises or consists of an antigen binding fragment selected from the group consisting of FAB-like molecules, such as Fab and F(ab')2, Fv molecules, disulphide-linked Fv molecules, ScFv molecules and single domain antibodies (dAbs).
- 16. (previously presented) A compound according to Claim 1 wherein the target specific portion comprises one or more antibody constant regions.
- 17. (original) A compound according to Claim 16 wherein the one or more antibody constant regions comprises or consists of a CH1 domain.
- 18. (previously presented) A compound according to Claim 1 further comprising an Fe moiety.
- 19. (original) A compound according to Claim 18 wherein the Fe moiety is derived from human IgG1.
- 20. (previously presented) A compound according to-Claim 1 wherein the target specific portion comprises or consists of a whole BC1 antibody.
- 21. (previously presented) A compound according to Claim 1 wherein the effector portion comprises or consists of human interleukin-12, or a functional fragment or variant thereof.
- 22. (previously presented) A compound according to Claim 1 wherein the effector portion comprises or consists of a single-chain interleukin-12.

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- 23. (previously presented) A compound according to Claim 22 wherein the single chain IL-12 consists of an Il-12p35 domain and an IL-12p40 domain.
- 24. (previously presented) A compound according to Claim 23 wherein the IL-12p35 domain is conjugated to the IL-12p40 domain by a disulphide bond.
- 25. (previously presented) A compound according to Claim 1 wherein the compound is a fusion protein.
- 26. (previously presented) A compound according to Claim 1 wherein the target specific portion is fused to the effector portion.
- 27. (original) A compound according to Claim 26 comprising an immunoglobulin heavy chain fused to the effector portion.
- 28. (original) A compound according to Claim 27 wherein the immunoglobulin heavy chain and the effector portion are joined via a mutated linker sequence.
- 29. (original) A compound according to Claim 28 wherein the linker comprises or consists of the amino acid sequence ATATPGAA (SEQ ID NO: 5).
- 30. (previously presented) A compound according to Claim 1 wherein the compound comprises a polypeptide of SEQ ID NO: 6.
- 31. (previously presented) A compound according to Claim 1 wherein the compound comprises a polypeptide of SEQ ID NO: 7.
- 32. (previously presented) A compound according to Claim 30 wherein the compound comprises a polypeptide of SEQ ID NO:6 and a polypeptide of SEQ ID NO:7.

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- 33. (previously presented) A compound according to Claim 30 further comprising a polypeptide of SEQ ID 4 linked by disulphide bond to the polypeptide of SEQ ID NO:6.
- 34. (original) A fusion protein comprising antibody V regions directed against oncofoetal fibronectin, an Fe moiety, and an interleukin-12 moiety.
 - 35.- 42. (canceled)
- 43. (previously presented) A pharmaceutical composition comprising a compound according to Claim 1 and a pharamaceutically acceptable carrier.
- 44. (original) A pharmaceutical composition according to Claim 43 wherein the composition is suitable for parenteral administration.
 - 45. (previously presented) A compound according to Claim 1 for use in medicine.
 - 46. (canceled)
- 47. (previously presented) A method of treating a patient with cancer, the method comprising administering a compound according to Claim 1 to said patient.
- 48. (previously presented) The method according to Claim 47 wherein the mammal is a human.
- 49. (previously presented) The method according to Claim 47 wherein the patient has a solid tumor.
- 50. (previously presented) The method according to Claim 47 wherein the cancer is a glioblastoma.
 - 51. (canceled)

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52. (canceled)